This Page Is Inserted by IFW Operations and is not a part of the Official Record

BEST AVAILABLE IMAGES

Defective images within this document are accurate representations of the original documents submitted by the applicant.

Defects in the images may include (but are not limited to):

- BLACK BORDERS
- TEXT CUT OFF AT TOP, BOTTOM OR SIDES
- FADED TEXT
- ILLEGIBLE TEXT
- SKEWED/SLANTED IMAGES
- COLORED PHOTOS
- BLACK OR VERY BLACK AND WHITE DARK PHOTOS
- GRAY SCALE DOCUMENTS

IMAGES ARE BEST AVAILABLE COPY.

As rescanning documents will not correct images, please do not report the images to the Image Problem Mailbox.



<211> 32

SEQUENCE LISTING

```
<110> Aventis Behring GmbH
<120> Modified cDNA Factor VIII and its Derivatives
<130> 2002/M018-A66
<140> US/10/721,997
<141> 2003-11-26
<160> 34
<170> PatentIn version 3.1
<210> 1
<211> 34
<212> DNA
<213> Artificial Sequence
<220>
<223> Human Factor VIII derivatives
<220>
<221> Forward primer
 <222> (1)..(34)
 <223>
 <400>
 ggaaccatcg ccagaagtcc ttggaaatct cgcc
 34
        2
 <210>
 <211> 34
 <212> DNA
 <213> Artificial Sequence
 <220>
 <223> Human Factor VIII derivatives
 <220>
 <221> Reverse primer
       (1)..(34)
 <222>
 <223>
 <400> 2
 ggcgagattt ccaaggactt ctggcgatgg ttcc
 34
 <210>
        3
```

1.1

```
<212> DNA
<213> Artificial Sequence
<220>
<223> Human Factor VIII derivatives
<220>
<221> Forward primer
<222> (1)..(32)
<223>
<400> 3
cccaccaaca tggtggcatg gaagcttatg tc
32
<210> 4
<211> 32
<212> DNA
<213> Artificial Sequence
 <220>
<223> Human Factor VIII derivatives
 <220>
 <221> Reverse primer
 <222> (1)..(32)
 <223>
 <400> 4
 gacataagct tccatgccac catgttggtg gg
 32
 <210> 5
 <211> 47
 <212> DNA
 <213> Artificial Sequence
 <220>
       Human Factor VIII derivatives
 <223>
 <220>
 <221> Forward primer
  <222> (1)..(47)
  <223>
  <400> 5
  cagaggaacc ccaactacga cgtaaaaata atgaagaagc ggaagac
  47
```

Z)

42

```
<210>
      47
<211>
<212> DNA
<213> Artificial Sequence
<220>
<223> Human Factor VIII derivatives
<220>
<221> Reverse primer
<222> (1)..(47)
<223>
<400> 6
gtetteeget tetteattat ttttaegteg tagttggggt teetetg
47
      7
 <210>
 <211> 42
 <212> DNA
 <213> Artificial Sequence
 <220>
 <223> Human Factor VIII derivatives
 <220>
 <221> Forward primer
 <222> (1)..(42)
 <223>
 <400> 7
 cccaactacg aatgaaaaat gatgaagaag cggaagacta tg
 <210> 8
 <211> 42
 <212> DNA
 <213> Artificial Sequence
 <220>
 <223> Human Factor VIII derivatives
 <220>
  <221> Reverse primer
  <222> (1)..(42)
  <223>
  catagtette egettettea teatttttea ttegtagttg gg
```

```
9
<210>
      40
<211>
<212> DNA
<213> Artificial Sequence
<220>
<223> Human Factor VIII derivatives
<220>
      Forward primer
<221>
<222> (1)..(40)
<223>
<400>
       9
gaagaagcgg aagactatga tgataatctt actgattctg
40
<210>
       10
       40
<211>
 <212> DNA
 <213> Artificial Sequence
 <220>
       Human Factor VIII derivatives
 <223>
 <220>
 <221> Reverse primer
       (1)..(40)
 <222>
 <223>
 <400>
        10
 cagaatcagt aagattatca tcatagtctt ccgcttcttc
 40
 <210>
        11
       38
 <211>
 <212> DNA
 <213> Artificial Sequence
 <220>
       Human Factor VIII derivatives
 <223>
 <220>
 <221> Forward primer
       (1)..(38)
 <222>
 <223>
```

<400> 11

```
ggtcaggttt gatgatgacg acteteette etttatee
38
<210> 12
<211> 38
<212> DNA
<213> Artificial Sequence
<220>
<223> Human Factor VIII derivatives
<220>
<221> Reverse primer
<222> (1)..(38)
<223>
 <400> 12
ggataaagga aggagagtcg tcatcatcaa acctgacc
 38
 <210> 13
 <211> 38
 <212> DNA
 <213> Artificial Sequence
 <220>
 <223> Human Factor VIII derivatives
 <220>
 <221> Forward primer
 <222>
       (1)..(38)
 <223>
 <400> 13
 cccttagtcc tcgcccctc tgacagaagt tataaaag
 38
 <210> 14
 <211> 38
 <212> DNA
 <213> Artificial Sequence
 <220>
 <223> Human Factor VIII derivatives
 <220>
 <221> Reverse primer
 <222> (1)..(38)
```

<223>

```
<400> 14
cttttataac ttctgtcaga gggggcgagg actaaggg
<210> 15
<211> 41
<212> DNA
<213> Artificial Sequence
<220>
       Human Factor VIII derivatives
<223>
<400> 15
gtccgattta tggcatacac agatgttacc tttaagactc g
41
<210> 16
<211> 41
<212> DNA
<213> Artificial Sequence
<220>
       Human Factor VIII derivatives
<223>
<220>
<221> Reverse primer
       (1)..(41)
 <222>
 <223>
 <400> 16
 cgagtcttaa aggtaacatc tgtgtatgcc ataaatcgga c
 41
 <210> 17
 <211> 37
 <212> DNA
 <213> Artificial Sequence
 <220>
 <223> Human Factor VIII derivatives
 <220>
 <221> Forward primer
       (1)..(37)
 <222>
 <223>
 <400>
 cctttaagac tcgtaaagct attcagcatg aatcagg
 37
```

Li

<400> 20

```
<210> 18
<211> 37
<212> DNA
<213> Artificial Sequence
<220>
<223> Human Factor VIII derivatives
<220>
<221> Reverse primer
<222> (1)..(37)
<223>
<400> 18
cctgattcat gctgaatagc tttacgagtc ttaaagg
37
<210> 19
<211> 46
<212> DNA
<213> Artificial Sequence
<220>
<223> Human Factor VIII derivatives
 <220>
 <221> Forward primer
 <222> (1)..(46)
 <223>
 <400> 19
 cacactgttg attatattta agaataaagc aagcagacca tataac
 <210> 20
 <211> 46
 <212> DNA
 <213> Artificial Sequence
 <220>
       Human Factor VIII derivatives
 <223>
 <220>
 <221> Reverse primer
       (1)..(46)
 <222>
 <223>
```

```
gttatatggt ctgcttgctt tattcttaaa tataatcaac agtgtg
46
<210> 21
<211> 40
<212> DNA
<213> Artificial Sequence
<220>
<223> Human Factor VIII derivatives
<220>
<221> Forward primer
<222> (1)..(40)
<223>
<400> 21
ccctcacgga atcactgatg tctctccttt gtattcaagg
40
<210> 22
<211> 40
<212> DNA
<213> Artificial Sequence
<220>
 <223> Human Factor VIII derivatives
 <220>
 <221> Reverse primer
       (1)..(40)
 <222>
 <223>
 <400> 22
 ccttgaatac aaaggagaga catcagtgat tccgtgaggg
 <210> 23
 <211> 38
 <212> DNA
 <213> Artificial Sequence
 <220>
 <223> Human Factor VIII derivatives
 <220>
 <221> Forward primer
 <222> (1)..(38)
```

<223>

```
<400>
      23
gatgtccgtc ctttgtattc agggagatta ccaaaagg
<210> 24
<211> 38
<212> DNA
<213> Artificial Sequence
<220>
      Human Factor VIII derivatives
<223'>
<220>
<221> Reverse primer
<222> (1)..(38)
 <223>
 <400> 24
ccttttggta atctccctga atacaaagga cggacatc
 38
 <210>
        25
 <211> 36
 <212> DNA
 <213> Artificial Sequence
 <220>
 <223> Human Factor VIII derivatives
 <220>
 <221> Forward primer
 <222> (1)..(36)
 <223> .
 <400> 25
 ctgtatttga tgagaaccaa agctggtacc tcacag
 36
        26
 <210>
 <211>
        36
 <212> DNA
 <213> Artificial Sequence
  <220>
 <223> Human Factor VIII derivatives
  <220>
  <221> Reverse primer
  <222> (1)..(36)
```

```
<223>
<400>
       26
ctgtgaggta ccagctttgg ttctcatcaa atacag
<210>
       27
       30
<211>
<212> DNA
<213> Artificial Sequence
<220>
       Human Factor VIII derivatives
<223>
<220>
<221> Forward primer
<222>
       (1)..(30)
<223>
<400>
       27
ctccccaatc cagatggagt gcagcttgag
30
<210>
       28
       30
<211>
<212> DNA
<213> Artificial Sequence
<220>
       Human Factor VIII derivatives
<223>
<220>
       Reverse primer
<221>
<222> (1)..(30)
<223>
<400> 28
ctcaagctgc actccatctg gattggggag
30
<210>
       29
 <211> 32
<212> DNA
<213> Artificial Sequence
<220>
       Human Factor VIII derivatives
 <223>
```

<220>

```
<221> Forward primer <222> (1)..(32)
<223>
<400> 29
cagetggagt geagetteag gateeagagt te
32
<210> 30
<211> 32
<212> DNA
<213> Artificial Sequence
<220>
<223> Human Factor VIII derivatives
<220>
<221> Reverse primer
<222> (1)..(32)
 <223>
 <400> 30
 gaactetgga teetgaaget geactecage tg
 <210> 31
 <211> 49
 <212> DNA
 <213> Artificial Sequence
 <220>
 <223> Human Factor VIII derivatives
 <220>
 <221> Forward primer
 <222> (1)..(49)
 <223>
 cgatggtatc tgctcagcat gaagagcaat gaaaacatcc attctattc
  <210> 32
 <211> 49
 <212> DNA
  <213> Artificial Sequence
  <220>
  <223> Human Factor VIII derivatives
```

```
<220>
```

<221> Reverse primer

<222> (1)..(49)

<223>

<400> 32 gaatagaatg gatgttttca ttgctcttca tgctgagcag ataccatcg 49

<210> 33

<211> 2114

<212> PRT

<213> Porcine

<400> 33

Ala Ile Arg Arg Tyr Tyr Leu Gly Ala Val Glu Leu Ser Trp Asp Tyr 1 5 10 15

Arg Gln Ser Glu Leu Leu Arg Glu Leu His Val Asp Thr Arg Phe Pro 20 25 30

Ala Thr Ala Pro Gly Ala Leu Pro Leu Gly Pro Ser Val Leu Tyr Lys 35 40 45

Lys Thr Val Phe Val Glu Phe Thr Asp Gln Leu Phe Ser Val Ala Arg 50 55 60

Pro Arg Pro Pro Trp Met Gly Leu Leu Gly Pro Thr Ile Gln Ala Glu 65 70 75 80

Val Tyr Asp Thr Val Val Val Thr Leu Lys Asn Met Ala Ser His Pro 85 90 95

Val Ser Leu His Ala Val Gly Val Ser Phe Trp Lys Ser Ser Glu Gly
100 105 110

Ala Glu Tyr Glu Asp His Thr Ser Gln Arg Glu Lys Glu Asp Asp Lys 115 120 125

Val Leu Pro Gly Lys Ser Gln Thr Tyr Val Trp Gln Val Leu Lys Glu 130 135 140 Asn Gly Pro Thr Ala Ser Asp Pro Pro Cys Leu Thr Tyr Ser Tyr Leu 145 150 155 160

Ser His Val Asp Leu Val Lys Asp Leu Asn Ser Gly Leu Ile Gly Ala 165 170 175

Leu Leu Val Cys Arg Glu Gly Ser Leu Thr Arg Glu Arg Thr Gln Asn 180 185 190

Leu His Glu Phe Val Leu Leu Phe Ala Val Phe Asp Glu Gly Lys Ser 195 200 205

Trp His Ser Ala Arg Asn Asp Ser Trp Thr Arg Ala Met Asp Pro Ala 210 215 220

Pro Ala Arg Ala Gln Pro Ala Met His Thr Val Asn Gly Tyr Val Asn 225 230 235 240

Arg Ser Leu Pro Gly Leu Ile Gly Cys His Lys Lys Ser Val Tyr Trp 245 250 255

His Val Ile Gly Met Gly Thr Ser Pro Glu Val His Ser Ile Phe Leu 260 265 270

Glu Gly His Thr Phe Leu Val Arg His His Arg Gln Ala Ser Leu Glu 275 280 285

Ile Ser Pro Leu Thr Phe Leu Thr Ala Gln Thr Phe Leu Met Asp Leu 290 295 300

Gly Gln Phe Leu Leu Phe Cys His Ile Ser Ser His His Gly Gly 305 310 315 320

Met Glu Ala His Val Arg Val Glu Ser Cys Ala Glu Glu Pro Gln Leu 325 330 335

Arg Arg Lys Ala Asp Glu Glu Glu Asp Tyr Asp Asp Asn Leu Tyr Asp 340 345 350

Ser Asp Met Asp Val Val Arg Leu Asp Gly Asp Asp Val Ser Pro Phe 355 360 365

Ile Gln Ile Arg Ser Val Ala Lys Lys His Pro Lys Thr Trp Val His 370 375 380

Tyr Ile Ser Ala Glu Glu Glu Asp Trp Asp Tyr Ala Pro Ala Val Pro 385 390 395 400

Ser Pro Ser Asp Arg Ser Tyr Lys Ser Leu Tyr Leu Asn Ser Gly Pro
405 410 415

Gln Arg Ile Gly Arg Lys Tyr Lys Lys Ala Arg Phe Val Ala Tyr Thr 420 425 430

Asp Val Thr Phe Lys Thr Arg Lys Ala Ile Pro Tyr Glu Ser Gly Ile 435 440 445

Leu Gly Pro Leu Leu Tyr Gly Glu Val Gly Asp Thr Leu Leu Ile Ile 450 455 460

Phe Lys Asn Lys Ala Ser Arg Pro Tyr Asn Ile Tyr Pro His Gly Ile 465 470 475 480

Thr Asp Val Ser Ala Leu His Pro Gly Arg Leu Leu Lys Gly Trp Lys 485 490 495

His Leu Lys Asp Met Pro Ile Leu Pro Gly Glu Thr Phe Lys Tyr Lys 500 505 510

Trp Thr Val Thr Val Glu Asp Gly Pro Thr Lys Ser Asp Pro Arg Cys 515 520 525

Leu Thr Arg Tyr Tyr Ser Ser Ser Ile Asn Leu Glu Lys Asp Leu Ala 530 540

Ser Gly Leu Ile Gly Pro Leu Leu Ile Cys Tyr Lys Glu Ser Val Asp 545 550 555 560

Gln Arg Gly Asn Gln Met Met Ser Asp Lys Arg Asn Val Ile Leu Phe 565 570 575

Ser Val Phe Asp Glu Asn Gln Ser Trp Tyr Leu Ala Glu Asn Ile Gln 580 585 590

Arg Phe Leu Pro Asn Pro Asp Gly Leu Gln Pro Gln Asp Pro Glu Phe 595 600 605

Gln Ala Ser Asn Ile Met His Ser Ile Asn Gly Tyr Val Phe Asp Ser 610 615 620

Leu Gln Leu Ser Val Cys Leu His Glu Val Ala Tyr Trp Tyr Ile Leu 625 630 635 640

Ser Val Gly Ala Gln Thr Asp Phe Leu Ser Val Phe Phe Ser Gly Tyr 645 650 655

Thr Phe Lys His Lys Met Val Tyr Glu Asp Thr Leu Thr Leu Phe Pro 660 665 670

Phe Ser Gly Glu Thr Val Phe Met Ser Met Glu Asn Pro Gly Leu Trp 675 680 685

Val Leu Gly Cys His Asn Ser Asp Leu Arg Asn Arg Gly Met Thr Ala 690 695 700

Leu Leu Lys Val Tyr Ser Cys Asp Arg Asp Ile Gly Asp Tyr Tyr Asp 705 710 715 720

Asn Thr Tyr Glu Asp Ile Pro Gly Phe Leu Leu Ser Gly Lys Asn Val 725 730 735

Ile Glu Pro Arg Ser Phe Ala Gln Asn Ser Arg Pro Pro Ser Ala Ser 740 745 750

Gln Lys Gln Phe Gln Thr Ile Thr Ser Pro Glu Asp Asp Val Glu Leu 755 760 765

Asp Pro Gln Ser Gly Glu Arg Thr Gln Ala Leu Glu Glu Leu Ser Val 770 775 780

Pro Ser Gly Asp Gly Ser Met Leu Leu Gly Gln Asn Pro Ala Pro His 785 790 795 800

Gly Ser Ser Ser Ser Asp Leu Gln Glu Ala Arg Asn Glu Ala Asp Asp

805 810 815

Tyr Leu Pro Gly Ala Arg Glu Arg Asn Thr Ala Pro Ser Ala Ala Ala 820 825 830

Arg Leu Arg Pro Glu Leu His His Ser Ala Glu Arg Val Leu Thr Pro 835 840 845

Glu Pro Glu Lys Glu Leu Lys Lys Leu Asp Ser Lys Met Ser Ser Ser 850 855

Ser Asp Leu Leu Lys Thr Ser Pro Thr Ile Pro Ser Asp Thr Leu Ser 865 870 875 880

Ala Glu Thr Glu Arg Thr His Ser Leu Gly Pro Pro His Pro Gln Val 885 890 895

Asn Phe Arg Ser Gln Leu Gly Ala Ile Val Leu Gly Lys Asn Ser Ser 900 905 910

His Phe Ile Gly Ala Gly Val Pro Leu Gly Ser Thr Glu Glu Asp His 915 920 925

Glu Ser Ser Leu Gly Glu Asn Val Ser Pro Val Glu Ser Asp Gly Ile 930 935 940

Phe Glu Lys Glu Arg Ala His Gly Pro Ala Ser Leu Thr Lys Asp Asp 945 950 955 960

Val Leu Phe Lys Val Asn Ile Ser Leu Val Lys Thr Asn Lys Ala Arg 965 970 975

Val Tyr Leu Lys Thr Asn Arg Lys Ile His Ile Asp Asp Ala Ala Leu 980 985 990

Leu Thr Glu Asn Arg Ala Ser Ala Thr Phe Met Asp Lys Asn Thr Thr 995 1000 1005

Ala Ser Gly Leu Asn His Val Ser Asn Trp Ile Lys Gly Pro Leu 1010 1015 1020

- Gly Lys Asn Pro Leu Ser Ser Glu Arg Gly Pro Ser Pro Glu Leu 1025 1030 1035
- Leu Thr Ser Ser Gly Ser Gly Lys Ser Val Lys Gly Gln Ser Ser 1040 1045 1050
- Gly Gln Gly Arg Ile Arg Val Ala Val Glu Glu Glu Leu Ser 1055 1060 1065
- Lys Gly Lys Glu Met Met Leu Pro Asn Ser Glu Leu Thr Phe Leu 1070 1075 1080
- Thr Asn Ser Ala Asp Val Gln Gly Asn Asp Thr His Ser Gln Gly 1085 1090 . 1095
- Lys Lys Ser Arg Glu Glu Met Glu Arg Arg Glu Lys Leu Val Gln 1100 1105 1110
- Glu Lys Val Asp Leu Pro Gln Val Tyr Thr Ala Thr Gly Thr Lys 1115 1120 1125
- Asn Phe Leu Arg Asn Ile Phe His Gln Ser Thr Glu Pro Ser Val 1130 1135 1140
- Glu Gly Phe Asp Gly Gly Ser His Ala Pro Val Pro Gln Asp Ser 1145 1150 1155
- Arg Ser Leu Asn Asp Ser Ala Glu Arg Ala Glu Thr His Ile Ala 1160 1165 1170
- His Phe Ser Ala Ile Arg Glu Glu Ala Pro Leu Glu Ala Pro Gly 1175 1180 1185
- Asn Arg Thr Gly Pro Gly Pro Arg Ser Ala Val Pro Arg Arg Val 1190 1195 1200
- Lys Gln Ser Leu Lys Gln Ile Arg Leu Pro Leu Glu Glu Ile Lys 1205 1210 1215
- Pro Glu Arg Gly Val Val Leu Asn Ala Thr Ser Thr Arg Trp Ser 1220 1225 1230

- Glu Ser Ser Pro Ile Leu Gln Gly Ala Lys Arg Asn Asn Leu Ser 1235 1240 1245
- Leu Pro Phe Leu Thr Leu Glu Met Ala Gly Gly Gln Gly Lys Ile 1250 1255 1260
- Ser Ala Leu Gly Lys Ser Ala Ala Gly Pro Leu Ala Ser Gly Lys 1265 1270 1275
- Leu Glu Lys Ala Val Leu Ser Ser Ala Gly Leu Ser Glu Ala Ser 1280 1285 1290
- Gly Lys Ala Glu Phe Leu Pro Lys Val Arg Val His Arg Glu Asp 1295 1300 1305
- Leu Leu Pro Gln Lys Thr Ser Asn Val Ser Cys Ala His Gly Asp 1310 1315 1320
- Leu Gly Gln Glu Ile Phe Leu Gln Lys Thr Arg Gly Pro Val Asn 1325 1330 1335
- Leu Asn Lys Val Asn Arg Pro Gly Arg Thr Pro Ser Lys Leu Leu 1340 1345 1350
- Gly Pro Pro Met Pro Lys Glu Trp Glu Ser Leu Glu Lys Ser Pro 1355 1360 1365
- Lys Ser Thr Ala Leu Arg Thr Lys Asp Ile Ile Ser Leu Pro Leu 1370 1375 1380
- Asp Arg His Glu Ser Asn His Ser Ile Ala Ala Lys Asn Glu Gly 1385 1390 1395
- Gln Ala Glu Thr Gln Arg Glu Ala Ala Trp Thr Lys Gln Gly Gly 1400 1405 1410
- Pro Gly Arg Leu Cys Ala Pro Lys Pro Pro Val Leu Arg Arg His 1415 1420 1425
- Gln Arg Asp Ile Ser Leu Pro Thr Phe Gln Pro Glu Glu Asp Lys 1430 1435 1440

- Met Asp Tyr Asp Asp Ile Phe Ser Thr Glu Thr Lys Gly Glu Asp 1445 1450 1455
- Phe Asp Ile Tyr Gly Glu Asp Glu Asn Gln Asp Pro Arg Ser Phe 1460 1465 1470
- Gln Lys Arg Thr Arg His Tyr Phe Ile Ala Ala Val Glu Gln Leu 1475 1480 1485
- Trp Asp Tyr Gly Met Ser Glu Ser Pro Arg Ala Leu Arg Asn Arg 1490 1495 1500
- Ala Gln Asn Gly Glu Val Pro Arg Phe Lys Lys Val Val Phe Arg 1505 1510 1515
- Glu Phe Ala Asp Gly Ser Phe Thr Gln Pro Ser Tyr Arg Gly Glu 1520 1525 1530
- Leu Asn Lys His Leu Gly Leu Leu Gly Pro Tyr Ile Arg Ala Glu 1535 1540 1545
- Val Glu Asp Asn Ile Met Val Thr Phe Lys Asn Gln Ala Ser Arg 1550 1555 1560
- Pro Tyr Ser Phe Tyr Ser Ser Leu Ile Ser Tyr Pro Asp Asp Gln 1565 1570 1575
- Glu Gln Gly Ala Glu Pro Arg His Asn Phe Val Gln Pro Asn Glu 1580 1585 1590
- Thr Arg Thr Tyr Phe Trp Lys Val Gln His His Met Ala Pro Thr 1595 1600 1605
- Glu Asp Glu Phe Asp Cys Lys Ala Trp Ala Tyr Phe Ser Asp Val 1610 1615 1620
- Asp Leu Glu Lys Asp Val His Ser Gly Leu Ile Gly Pro Leu Leu 1625 1630 1635
- Ile Cys Arg Ala Asn Thr Leu Asn Ala Ala His Gly Arg Gln Val

1640 1645 1650

Thr Val Gln Glu Phe Ala Leu Phe Phe Thr Ile Phe Asp Glu Thr 1655 1660 1665

- Lys Ser Trp Tyr Phe Thr Glu Asn Val Glu Arg Asn Cys Arg Ala 1670 1675 1680
- Pro Cys His Leu Gln Met Glu Asp Pro Thr Leu Lys Glu Asn Tyr 1685 1690 1695
- Arg Phe His Ala Ile Asn Gly Tyr Val Met Asp Thr Leu Pro Gly 1700 1705 1710
- Leu Val Met Ala Gln Asn Gln Arg Ile Arg Trp Tyr Leu Leu Ser 1715 1720 1725
- Met Gly Ser Asn Glu Asn Ile His Ser Ile His Phe Ser Gly His 1730 1735 1740
- Val Phe Ser Val Arg Lys Lys Glu Glu Tyr Lys Met Ala Val Tyr 1745 1750 1755
- Asn Leu Tyr Pro Gly Val Phe Glu Thr Val Glu Met Leu Pro Ser 1760 1765 1770
- Lys Val Gly Ile Trp Arg Ile Glu Cys Leu Ile Gly Glu His Leu 1775 1780 1785
- Gln Ala Gly Met Ser Thr Thr Phe Leu Val Tyr Ser Lys Glu Cys 1790 1795 1800
- Gln Ala Pro Leu Gly Met Ala Ser Gly Arg Ile Arg Asp Phe Gln 1805 1810 1815
- Ile Thr Ala Ser Gly Gln Tyr Gly Gln Trp Ala Pro Lys Leu Ala 1820 1825 1830
- Arg Leu His Tyr Ser Gly Ser Ile Asn Ala Trp Ser Thr Lys Asp 1835 1840 1845

- Pro His Ser Trp Ile Lys Val Asp Leu Leu Ala Pro Met Ile Ile 1850 1855 1860
- His Gly Ile Met Thr Gln Gly Ala Arg Gln Lys Phe Ser Ser Leu 1865 1870 1875
- Tyr Ile Ser Gln Phe Ile Ile Met Tyr Ser Leu Asp Gly Arg Asn 1880 1885 1890
- Trp Gln Ser Tyr Arg Gly Asn Ser Thr Gly Thr Leu Met Val Phe 1895 1900 1905
- Phe Gly Asn Val Asp Ala Ser Gly Ile Lys His Asn Ile Phe Asn 1910 1915 1920
- Pro Pro Ile Val Ala Arg Tyr Ile Arg Leu His Pro Thr His Tyr 1925 1930 1935
 - Ser Ile Arg Ser Thr Leu Arg Met Glu Leu Met Gly Cys Asp Leu 1940 1945 1950
 - Asn Ser Cys Ser Met Pro Leu Gly Met Gln Asn Lys Ala Ile Ser 1955 1960 1965
 - Asp Ser Gln Ile Thr Ala Ser Ser His Leu Ser Asn Ile Phe Ala 1970 1975 1980
 - Thr Trp Ser Pro Ser Gln Ala Arg Leu His Leu Gln Gly Arg Thr 1985 1990 1995
 - Asn Ala Trp Arg Pro Arg Val Ser Ser Ala Glu Glu Trp Leu Gln
 2000 2005 2010
 - Val Asp Leu Gln Lys Thr Val Lys Val Thr Gly Ile Thr Thr Gln 2015 2020 2025
 - Gly Val Lys Ser Leu Leu Ser Ser Met Tyr Val Lys Glu Phe Leu 2030 2035 2040
 - Val Ser Ser Ser Gln Asp Gly Arg Arg Trp Thr Leu Phe Leu Gln 2045 2050 2055

Asp Gly His Thr Lys Val Phe Gln Gly Asn Gln Asp Ser Ser Thr 2060 2065 2070

Pro Val Val Asn Ala Leu Asp Pro Pro Leu Phe Thr Arg Tyr Leu 2075 2080 2085

Arg Ile His Pro Thr Ser Trp Ala Gln His Ile Ala Leu Arg Leu 2090 2095 2100

Glu Val Leu Gly Cys Glu Ala Gln Asp Leu Tyr 2105 2110

<210> 34

<211> 2332

<212> PRT

<213> Homo sapiens

<400> 34

Ala Thr Arg Arg Tyr Tyr Leu Gly Ala Val Glu Leu Ser Trp Asp Tyr 1 5 10 15

Met Gln Ser Asp Leu Gly Glu Leu Pro Val Asp Ala Arg Phe Pro Pro 20 25 30

Arg Val Pro Lys Ser Phe Pro Phe Asn Thr Ser Val Val Tyr Lys Lys 35 40 45

Thr Leu Phe Val Glu Phe Thr Asp His Leu Phe Asn Ile Ala Lys Pro 50 55 60

Arg Pro Pro Trp Met Gly Leu Leu Gly Pro Thr Ile Gln Ala Glu Val 65 70 75 80

Tyr Asp Thr Val Val Ile Thr Leu Lys Asn Met Ala Ser His Pro Val 85 90 95

Ser Leu His Ala Val Gly Val Ser Tyr Trp Lys Ala Ser Glu Gly Ala 100 105 110

Glu Tyr Asp Asp Gln Thr Ser Gln Arg Glu Lys Glu Asp Asp Lys Val 115 120 125 Phe Pro Gly Gly Ser His Thr Tyr Val Trp Gln Val Leu Lys Glu Asn 130 135 140

Gly Pro Met Ala Ser Asp Pro Leu Cys Leu Thr Tyr Ser Tyr Leu Ser 145 150 155 160

His Val Asp Leu Val Lys Asp Leu Asn Ser Gly Leu Ile Gly Ala Leu 165 170 175

Leu Val Cys Arg Glu Gly Ser Leu Ala Lys Glu Lys Thr Gln Thr Leu 180 185 190

His Lys Phe Ile Leu Leu Phe Ala Val Phe Asp Glu Gly Lys Ser Trp 195 200 205

His Ser Glu Thr Lys Asn Ser Leu Met Gln Asp Arg Asp Ala Ala Ser 210 215 220

Ala Arg Ala Trp Pro Lys Met His Thr Val Asn Gly Tyr Val Asn Arg 225 230 235 240

Ser Leu Pro Gly Leu Ile Gly Cys His Arg Lys Ser Val Tyr Trp His 245 250 255

Val Ile Gly Met Gly Thr Thr Pro Glu Val His Ser Ile Phe Leu Glu 260 265 270

Gly His Thr Phe Leu Val Arg Asn His Arg Gln Ala Ser Leu Glu Ile 275 280 285

Ser Pro Ile Thr Phe Leu Thr Ala Gln Thr Leu Leu Met Asp Leu Gly 290 295 300

Gln Phe Leu Leu Phe Cys His Ile Ser Ser His Gln His Asp Gly Met 305 310 315 320

Glu Ala Tyr Val Lys Val Asp Ser Cys Pro Glu Glu Pro Gln Leu Arg 325 330 335

Met Lys Asn Asn Glu Glu Ala Glu Asp Tyr Asp Asp Asp Leu Thr Asp 340 345 350

Ser Glu Met Asp Val Val Arg Phe Asp Asp Asp Asn Ser Pro Ser Phe 355 360 365

10

Ile Gln Ile Arg Ser Val Ala Lys Lys His Pro Lys Thr Trp Val His 370 375 380

Tyr Ile Ala Ala Glu Glu Glu Asp Trp Asp Tyr Ala Pro Leu Val Leu 385 390 395 400

Ala Pro Asp Asp Arg Ser Tyr Lys Ser Gln Tyr Leu Asn Asn Gly Pro 405 410 415

Gln Arg Ile Gly Arg Lys Tyr Lys Lys Val Arg Phe Met Ala Tyr Thr 420 425 430

Asp Glu Thr Phe Lys Thr Arg Glu Ala Ile Gln His Glu Ser Gly Ile 435 440 445

Leu Gly Pro Leu Leu Tyr Gly Glu Val Gly Asp Thr Leu Leu Ile Ile 450 455 460

Phe Lys Asn Gln Ala Ser Arg Pro Tyr Asn Ile Tyr Pro His Gly Ile 465 470 475 480

Thr Asp Val Arg Pro Leu Tyr Ser Arg Arg Leu Pro Lys Gly Val Lys
485 490 495

His Leu Lys Asp Phe Pro Ile Leu Pro Gly Glu Ile Phe Lys Tyr Lys 500 505 510

Trp Thr Val Thr Val Glu Asp Gly Pro Thr Lys Ser Asp Pro Arg Cys 515 520 525

Leu Thr Arg Tyr Tyr Ser Ser Phe Val Asn Met Glu Arg Asp Leu Ala 530 535 540

Ser Gly Leu Ile Gly Pro Leu Leu Ile Cys Tyr Lys Glu Ser Val Asp 545 550 555 560

Gln Arg Gly Asn Gln Ile Met Ser Asp Lys Arg Asn Val Ile Leu Phe

565 570 575

T₁, •

Ser Val Phe Asp Glu Asn Arg Ser Trp Tyr Leu Thr Glu Asn Ile Gln Arg Phe Leu Pro Asn Pro Ala Gly Val Gln Leu Glu Asp Pro Glu Phe Gln Ala Ser Asn Ile Met His Ser Ile Asn Gly Tyr Val Phe Asp Ser Leu Gln Leu Ser Val Cys Leu His Glu Val Ala Tyr Trp Tyr Ile Leu Ser Ile Gly Ala Gln Thr Asp Phe Leu Ser Val Phe Phe Ser Gly Tyr Thr Phe Lys His Lys Met Val Tyr Glu Asp Thr Leu Thr Leu Phe Pro Phe Ser Gly Glu Thr Val Phe Met Ser Met Glu Asn Pro Gly Leu Trp Ile Leu Gly Cys His Asn Ser Asp Phe Arg Asn Arg Gly Met Thr Ala Leu Leu Lys Val Ser Ser Cys Asp Lys Asn Thr Gly Asp Tyr Tyr Glu Asp Ser Tyr Glu Asp Ile Ser Ala Tyr Leu Leu Ser Lys Asn Asn Ala Ile Glu Pro Arg Ser Phe Ser Gln Asn Ser Arg His Arg Ser Thr Arg

Gln Lys Gln Phe Asn Ala Thr Thr Ile Pro Glu Asn Asp Ile Glu Lys

Thr Asp Pro Trp Phe Ala His Arg Thr Pro Met Pro Lys Ile Gln Asn 770 775 780

Val Ser Ser Ser Asp Leu Leu Met Leu Leu Arg Gln Ser Pro Thr Pro 785 790 795 800

His Gly Leu Ser Leu Ser Asp Leu Gln Glu Ala Lys Tyr Glu Thr Phe 805 810 815

Ser Asp Asp Pro Ser Pro Gly Ala Ile Asp Ser Asn Asn Ser Leu Ser 820 825 830

Glu Met Thr His Phe Arg Pro Gln Leu His His Ser Gly Asp Met Val 835 840 845

Phe Thr Pro Glu Ser Gly Leu Gln Leu Arg Leu Asn Glu Lys Leu Gly 850 855 860

Thr Thr Ala Ala Thr Glu Leu Lys Lys Leu Asp Phe Lys Val Ser Ser 865 870 875 880

Thr Ser Asn Asn Leu Ile Ser Thr Ile Pro Ser Asp Asn Leu Ala Ala 885 890 895

Gly Thr Asp Asn Thr Ser Ser Leu Gly Pro Pro Ser Met Pro Val His
900 905 910

Tyr Asp Ser Gln Leu Asp Thr Thr Leu Phe Gly Lys Lys Ser Ser Pro 915 920 925

Leu Thr Glu Ser Gly Gly Pro Leu Ser Leu Ser Glu Glu Asn Asn Asp 930 935 940

Ser Lys Leu Leu Glu Ser Gly Leu Met Asn Ser Gln Glu Ser Ser Trp 945 950 955 960

Gly Lys Asn Val Ser Ser Thr Glu Ser Gly Arg Leu Phe Lys Gly Lys 965 970 975

Arg Ala His Gly Pro Ala Leu Leu Thr Lys Asp Asn Ala Leu Phe Lys 980 985 990

Val Ser Ile Ser Leu Leu Lys Thr Asn Lys Thr Ser Asn Asn Ser Ala 995 1000 1005

- Thr Asn Arg Lys Thr His Ile Asp Gly Pro Ser Leu Leu Ile Glu 1010 1015 1020
- Asn Ser Pro Ser Val Trp Gln Asn Ile Leu Glu Ser Asp Thr Glu 1025 1030 1035
- Phe Lys Lys Val Thr Pro Leu Ile His Asp Arg Met Leu Met Asp 1040 1045 1050
- Lys Asn Ala Thr Ala Leu Arg Leu Asn His Met Ser Asn Lys Thr 1055 1060 1065
- Thr Ser Ser Lys Asn Met Glu Met Val Gln Gln Lys Lys Glu Gly
 1070 1075 1080
- Pro Ile Pro Pro Asp Ala Gln Asn Pro Asp Met Ser Phe Phe Lys 1085 1090 1095
- Met Leu Phe Leu Pro Glu Ser Ala Arg Trp Ile Gln Arg Thr His 1100 1105 1110
- Gly Lys Asn Ser Leu Asn Ser Gly Gln Gly Pro Ser Pro Lys Gln 1115 1120 1125
- Leu Val Ser Leu Gly Pro Glu Lys Ser Val Glu Gly Gln Asn Phe 1130 1135 1140
- Leu Ser Glu Lys Asn Lys Val Val Gly Lys Gly Glu Phe Thr 1145 1150 1155
- Lys Asp Val Gly Leu Lys Glu Met Val Phe Pro Ser Ser Arg Asn 1160 1165 1170
- Leu Phe Leu Thr Asn Leu Asp Asn Leu His Glu Asn Asn Thr His 1175 1180 1185
- Asn Gln Glu Lys Lys Ile Gln Glu Glu Ile Glu Lys Lys Glu Thr 1190 1195 1200
- Leu Ile Gln Glu Asn Val Val Leu Pro Gln Ile His Thr Val Thr 1205 1210 1215

- Gly Thr Lys Asn Phe Met Lys Asn Leu Phe Leu Leu Ser Thr Arg 1220 1225 1230
- Gln Asn Val Glu Gly Ser Tyr Asp Gly Ala Tyr Ala Pro Val Leu 1235 1240 1245
- Gln Asp Phe Arg Ser Leu Asn Asp Ser Thr Asn Arg Thr Lys Lys 1250 1255 1260
- His Thr Ala His Phe Ser Lys Lys Gly Glu Glu Glu Asn Leu Glu 1265 1270 1275
- Gly Leu Gly Asn Gln Thr Lys Gln Ile Val Glu Lys Tyr Ala Cys 1280 1285 1290
- Thr Thr Arg Ile Ser Pro Asn Thr Ser Gln Gln Asn Phe Val Thr 1295 1300 1305
- Gln Arg Ser Lys Arg Ala Leu Lys Gln Phe Arg Leu Pro Leu Glu 1310 1315 1320
- Glu Thr Glu Leu Glu Lys Arg Ile Ile Val Asp Asp Thr Ser Thr 1325 1330 1335
- Gln Trp Ser Lys Asn Met Lys His Leu Thr Pro Ser Thr Leu Thr 1340 1345 1350
- Gln Ile Asp Tyr Asn Glu Lys Glu Lys Gly Ala Ile Thr Gln Ser 1355 1360 1365
- Pro Leu Ser Asp Cys Leu Thr Arg Ser His Ser Ile Pro Gln Ala 1370 1375 1380
- Asn Arg Ser Pro Leu Pro Ile Ala Lys Val Ser Ser Phe Pro Ser 1385 1390 1395
- Ile Arg Pro Ile Tyr Leu Thr Arg Val Leu Phe Gln Asp Asn Ser 1400 1405 1410
- Ser His Leu Pro Ala Ala Ser Tyr Arg Lys Lys Asp Ser Gly Val

n , s

Gln Glu Ser Ser His Phe Leu Gln Gly Ala Lys Lys Asn Asn Leu Ser Leu Ala Ile Leu Thr Leu Glu Met Thr Gly Asp Gln Arg Glu Val Gly Ser Leu Gly Thr Ser Ala Thr Asn Ser Val Thr Tyr Lys Lys Val Glu Asn Thr Val Leu Pro Lys Pro Asp Leu Pro Lys Thr Ser Gly Lys Val Glu Leu Leu Pro Lys Val His Ile Tyr Gln Lys Asp Leu Phe Pro Thr Glu Thr Ser Asn Gly Ser Pro Gly His Leu Asp Leu Val Glu Gly Ser Leu Leu Gln Gly Thr Glu Gly Ala Ile Lys Trp Asn Glu Ala Asn Arg Pro Gly Lys Val Pro Phe Leu Arg Val Ala Thr Glu Ser Ser Ala Lys Thr Pro Ser Lys Leu Leu Asp Pro Leu Ala Trp Asp Asn His Tyr Gly Thr Gln Ile Pro Lys Glu Glu Trp Lys Ser Gln Glu Lys Ser Pro Glu Lys Thr Ala Phe Lys Lys Lys Asp Thr Ile Leu Ser Leu Asn Ala Cys Glu Ser Asn His

Ala Ile Ala Ile Asn Glu Gly Gln Asn Lys Pro Glu Ile Glu

1 5. o

Val Thr Trp Ala Lys Gln Gly Arg Thr Glu Arg Leu Cys Ser Gln Asn Pro Pro Val Leu Lys Arg His Gln Arg Glu Ile Thr Arg Thr Thr Leu Gln Ser Asp Gln Glu Glu Ile Asp Tyr Asp Asp Thr Ile Ser Val Glu Met Lys Lys Glu Asp Phe Asp Ile Tyr Asp Glu Asp Glu Asn Gln Ser Pro Arg Ser Phe Gln Lys Lys Thr Arg His Tyr Phe Ile Ala Ala Val Glu Arg Leu Trp Asp Tyr Gly Met Ser Ser Ser Pro His Val Leu Arg Asn Arg Ala Gln Ser Gly Ser Val Pro Gln Phe Lys Lys Val Val Phe Gln Glu Phe Thr Asp Gly Ser Phe Thr Gln Pro Leu Tyr Arg Gly Glu Leu Asn Glu His Leu Gly Leu Leu Gly Pro Tyr Ile Arq Ala Glu Val Glu Asp Asn Ile Met Val Thr Phe Arg Asn Gln Ala Ser Arg Pro Tyr Ser Phe Tyr Ser Ser Leu Ile Ser Tyr Glu Glu Asp Gln Arg Gln Gly Ala Glu Pro Arg Lys Asn Phe Val Lys Pro Asn Glu Thr Lys Thr Tyr Phe Trp Lys Val Gln His His Met Ala Pro Thr Lys Asp Glu Phe Asp Cys Lys

- Ala Trp Ala Tyr Phe Ser Asp Val Asp Leu Glu Lys Asp Val His 1835 1840 1845
- Ser Gly Leu Ile Gly Pro Leu Leu Val Cys His Thr Asn Thr Leu 1850 1855 1860
- Asn Pro Ala His Gly Arg Gln Val Thr Val Gln Glu Phe Ala Leu 1865 1870 1875
- Phe Phe Thr Ile Phe Asp Glu Thr Lys Ser Trp Tyr Phe Thr Glu 1880 1885 1890
- Asn Met Glu Arg Asn Cys Arg Ala Pro Cys Asn Ile Gln Met Glu 1895 1900 1905
- Asp Pro Thr Phe Lys Glu Asn Tyr Arg Phe His Ala Ile Asn Gly 1910 1915 1920
- Tyr Ile Met Asp Thr Leu Pro Gly Leu Val Met Ala Gln Asp Gln 1925 1930 1935
- Arg Ile Arg Trp Tyr Leu Leu Ser Met Gly Ser Asn Glu Asn Ile 1940 1945 1950
- His Ser Ile His Phe Ser Gly His Val Phe Thr Val Arg Lys Lys 1955 1960 1965
- Glu Glu Tyr Lys Met Ala Leu Tyr Asn Leu Tyr Pro Gly Val Phe 1970 1975 1980
- Glu Thr Val Glu Met Leu Pro Ser Lys Ala Gly Ile Trp Arg Val 1985 1990 1995
- Glu Cys Leu Ile Gly Glu His Leu His Ala Gly Met Ser Thr Leu 2000 2005 2010
- Phe Leu Val Tyr Ser Asn Lys Cys Gln Thr Pro Leu Gly Met Ala 2015 2020 2025
- Ser Gly His Ile Arg Asp Phe Gln Ile Thr Ala Ser Gly Gln Tyr 2030 2035 2040

Gly Gln Trp Ala Pro Lys Leu Ala Arg Leu His Tyr Ser Gly Ser 2045 2050 2055

4 .

- Ile Asn Ala Trp Ser Thr Lys Glu Pro Phe Ser Trp Ile Lys Val 2060 2065 2070
- Asp Leu Leu Ala Pro Met Ile Ile His Gly Ile Lys Thr Gln Gly 2075 2080 2085
- Ala Arg Gln Lys Phe Ser Ser Leu Tyr Ile Ser Gln Phe Ile Ile 2090 2095 2100
- Met Tyr Ser Leu Asp Gly Lys Lys Trp Gln Thr Tyr Arg Gly Asn 2105 2110 2115
- Ser Thr Gly Thr Leu Met Val Phe Phe Gly Asn Val Asp Ser Ser 2120 2125 2130
- Gly Ile Lys His Asn Ile Phe Asn Pro Pro Ile Ile Ala Arg Tyr 2135 2140 2145
- Ile Arg Leu His Pro Thr His Tyr Ser Ile Arg Ser Thr Leu Arg 2150 2155 2160
- Met Glu Leu Met Gly Cys Asp Leu Asn Ser Cys Ser Met Pro Leu 2165 2170 2175
- Gly Met Glu Ser Lys Ala Ile Ser Asp Ala Gln Ile Thr Ala Ser 2180 2185 2190
- Ser Tyr Phe Thr Asn Met Phe Ala Thr Trp Ser Pro Ser Lys Ala 2195 2200 2205
- Arg Leu His Leu Gln Gly Arg Ser Asn Ala Trp Arg Pro Gln Val 2210 2215 2220
- Asn Asn Pro Lys Glu Trp Leu Gln Val Asp Phe Gln Lys Thr Met 2225 2230 2235
- Lys Val Thr Gly Val Thr Thr Gln Gly Val Lys Ser Leu Leu Thr

2240 2245 2250

Ser Met Tyr Val Lys Glu Phe Leu Ile Ser Ser Ser Gln Asp Gly 2255 2260 2265

His Gln Trp Thr Leu Phe Phe Gln Asn Gly Lys Val Lys Val Phe 2270 2275 2280

Gln Gly Asn Gln Asp Ser Phe Thr Pro Val Val Asn Ser Leu Asp 2285 2290 2295

Pro Pro Leu Leu Thr Arg Tyr Leu Arg Ile His Pro Gln Ser Trp 2300 2305 2310

Val His Gln Ile Ala Leu Arg Met Glu Val Leu Gly Cys Glu Ala 2315 2320 2325

Gln Asp Leu Tyr 2330